## Amendments to the Claims:

Please amend claims 7 and 25-30, and add new claims 31-73. Claims 1-6 and 8-24 are canceled. This listing of claims will replace all prior versions, and listings of claims in the application:

## **Listing of Claims:**

- 1-6. (Canceled)
- 7. (Currently Amended): [[An]] <u>A</u> nucleic acid composition comprising: a nucleic acid vector having at least one cytosine to non-cytosine substitution within a CpG motif, wherein the CpG motif is of the formula 5' purine pyrimidine C G-pyrimidine pyrimidine 3' or 5' purine purine C G pyrimidine pyrimidine 3', and wherein the cytosine to non-cytosine substitution is within the CpG dinucleotide comprising a nucleic acid sequence of SEQ ID NO:297, wherein nucleotides at positions 784, 1161, 1218, 1264, 1337, 1829, 1831, 1874, 1876, 1940, 1942, 1963, 1966, 1987, 1997 and 1999 are as follows:

G at nucleotides 784, 1161, 1218, 1831, 1876, 1942, 1966 and 1999;

A at nucleotides 1264, 1337, 1829, 1874, 1940 and 1997; and

T at nucleotides 1963 and 1987.

- 8-24. (Canceled)
- 25. (Currently Amended): The nucleic acid composition of claim 7, wherein the nucleic acid vector composition further comprises an immune modulatory inhibitory nucleic acid sequence (IIS) comprising a hexamer region of the formula 5'-Purine-Purine-[X]-[Y]-Pyrimidine-Pyrimidine-3'; wherein X and Y are any naturally occurring or synthetic nucleotides except that X and Y cannot be cytosine-guanine.

- 26. (Currently Amended): The method nucleic acid composition of claim 25, wherein the immune modulatory nucleic acid IIS further comprises a polyG region linked 5' or 3' to the hexamer region.
- 27. (Currently Amended): The method nucleic acid composition of claim 25, wherein the immune modulatory nucleic acid IIS further comprises a first polyG region linked 5' to the hexamer region and a second polyG region linked 3' to the hexamer region.
- 28. (Currently Amended): The nucleic acid composition of claim 7, wherein the nucleic acid vector composition further comprises an immune modulatory nucleic acid IIS comprising a hexamer region of the formula 5'-Purine-Pyrimidine-[X]-[Y]-Pyrimidine-Pyrimidine-3', wherein X and Y are any naturally occurring or synthetic nucleotides except that X and Y cannot be cytosine-guanine.
- 29. (Currently Amended): The method nucleic acid composition of claim 28, wherein the immune modulatory nucleic acid IIS further comprises a polyG region linked 5' or 3' to the hexamer region.
- 30. (Currently Amended): The method nucleic acid composition of claim 28, wherein the immune modulatory nucleic acid <u>IIS</u> further comprises a first polyG region linked 5' to the hexamer region and a second polyG region linked 3' to the hexamer region.
- 31. (New): The nucleic acid composition of claim 25, wherein the nucleic acid vector further comprises the IIS.
- 32. (New): The nucleic acid composition of claim 7, wherein the vector further comprises a polynucleotide encoding an autoantigen targeted in an autoimmune disease.
- 33. (New): The nucleic acid composition of claim 32, wherein the autoantigen comprises a polynucleotide encoding a myelin protein.

- 34. (New): The nucleic acid composition of claim 33, wherein the myelin protein is myelin basic protein (MBP).
- 35. (New): The nucleic acid composition of claim 32, wherein the autoantigen comprises a polynucleotide encoding an insulin protein.
- 36. (New): The nucleic acid composition of claim 35, wherein the insulin protein is selected from the group consisting of insulin, proinsulin and preproinsulin.
- 37. (New): The nucleic acid composition of claim 7, further comprising a pharmaceutically acceptable carrier.
- 38. (New): A composition comprising a modified nucleic acid vector with reduced immunostimulatory properties, the nucleic acid vector modified by a method comprising the steps of:
- a) providing an unmodified nucleic acid vector comprising a CpG dinucleotide, wherein the CpG dinucleotide is in a motif of a formula 5'-purine-pyrimidine-C-G-pyrimidine-pyrimidine-3';
- b) substituting the cytosine in the CpG dinucleotide to a non-cytosine in the motif in the unmodified vector; thereby producing a modified nucleic acid vector, wherein the modified nucleic acid vector induces a reduced degree of immunostimulation in comparison to the unmodified nucleic acid vector.
- 39. (New): The composition of claim 38, wherein the cytosine to non-cytosine substitution is cytosine to guanine.
- 40. (New): The composition of claim 38, wherein a plurality of cytosine to non-cytosine substitutions are made.
- 41. (New): The composition of claim 40, wherein the plurality of cytosine to non-cytosine substitutions are made outside of a control region of the modified vector.

- 42. (New): The composition of claim 38, wherein the modified vector is a plasmid or cosmid vector.
- 43. (New): The composition of claim 38, wherein the composition further comprises an IIS comprising a hexamer region of a formula selected from the group consisting of 5'-Purine-Purine-[X]-[Y]-Pyrimidine-Pyrimidine-3' and 5'-Purine-Pyrimidine-[X]-[Y]-Pyrimidine-Pyrimidine-3'; wherein X and Y are any naturally occurring or synthetic nucleotides except that X and Y cannot be cytosine-guanine.
- 44. (New): The composition of claim 43, wherein the nucleic acid vector further comprises the IIS.
- 45. (New): The composition of claim 43, wherein the IIS further comprises a polyG region linked 5' or 3' to the hexamer region.
- 46. (New): The composition of claim 43, wherein the IIS further comprises a first polyG region linked 5' to the hexamer region and a second polyG region linked 3' to the hexamer region.
- 47. (New): The composition of claim 38, wherein the unmodified vector is SEQ ID NO:297.
- 48. (New): The composition of claim 47, wherein the unmodified vector that is SEQ ID NO:297 is modified to comprise the following cytosine to non-cytosine substitutions:

C to G at nucleotides 784, 1161, 1218 and 1966;

C to A at nucleotides 1264, 1337, 1829, 1874, 1940, and 1997; and

C to T at nucleotides 1963 and 1987.

49. (New): The composition of claim 48, wherein the unmodified vector that is SEQ ID NO:297 is further modified to comprise the following cytosine to non-cytosine substitutions: C to G at nucleotides 1831, 1876, 1942, and 1999.

- 50. (New): The composition of claim 38, further comprising a pharmaceutically acceptable carrier.
- 51. (New): The composition of claim 38, wherein the modified vector further comprises a polynucleotide encoding an autoantigen targeted in an autoimmune disease.
- 52. (New): The composition of claim 51, further comprising a polynucleotide encoding a myelin protein.
- 53. (New): The composition of claim 52, wherein the myelin protein is myelin basic protein (MBP).
- 54. (New): The composition of claim 51, further comprising a polynucleotide encoding an insulin protein.
- 55. (New): The composition of claim 54, wherein the insulin protein is selected from the group consisting of insulin, proinsulin and preproinsulin.
- 56. (New): A method of producing a modified nucleic acid vector with reduced immunostimulatory properties, the method comprising the steps of:
- a) providing an unmodified nucleic acid vector comprising a CpG dinucleotide, wherein the CpG dinucleotide is in a motif of a formula 5'-purine-pyrimidine-C-G-pyrimidine-pyrimidine-3';
- b) substituting the cytosine in the CpG dinucleotide to a non-cytosine in the motif in the unmodified vector; thereby producing a modified nucleic acid vector, wherein the modified nucleic acid vector induces a reduced degree of immunostimulation in comparison to the unmodified nucleic acid vector.
- 57. (New): The method of claim 56, wherein the cytosine to non-cytosine substitution is cytosine to guanine.
- 58. (New): The method of claim 56, wherein a plurality of cytosine to non-cytosine substitutions are made.

- 59. (New): The method of claim 58, wherein the plurality of cytosine to non-cytosine substitutions are made outside of a control region of the modified vector.
- 60. (New): The method of claim 56, wherein the modified vector is a plasmid or cosmid vector.
- 61. (New): The method of claim 56, wherein the composition further comprises an IIS comprising a hexamer region of a formula selected from the group consisting of 5'-Purine-Purine-[X]-[Y]-Pyrimidine-Pyrimidine-3' and 5'-Purine-Pyrimidine-[X]-[Y]-Pyrimidine-3'; wherein X and Y are any naturally occurring or synthetic nucleotides except that X and Y cannot be cytosine-guanine.
- 62. (New): The composition of claim 61, wherein the nucleic acid vector further comprises the IIS.
- 63. (New): The method of claim 61, wherein the IIS further comprises a polyG region linked 5' or 3' to the hexamer region.
- 64. (New): The method of claim 61, wherein the IIS further comprises a first polyG region linked 5' to the hexamer region and a second polyG region linked 3' to the hexamer region.
- 65. (New): The method of claim 56, wherein the unmodified vector is SEQ ID NO:297.
- 66. (New): The method of claim 65, wherein the unmodified vector that is SEQ ID NO:297 is modified to comprise the following cytosine to non-cytosine substitutions:

C to G at nucleotides 784, 1161, 1218 and 1966;

C to A at nucleotides 1264, 1337, 1829, 1874, 1940, and 1997; and

C to T at nucleotides 1963 and 1987.

- 67. (New): The method of claim 66, wherein the unmodified vector that is SEQ ID NO:297 is further modified to comprise the following cytosine to non-cytosine substitutions: C to G at nucleotides 1831, 1876, 1942, and 1999.
- 68. (New): The method of claim 56, further comprising a pharmaceutically acceptable carrier.
- 69. (New): The method of claim 56, wherein the modified vector further comprises a polynucleotide encoding an autoantigen targeted in an autoimmune disease.
- 70. (New): The method of claim 69, further comprising a polynucleotide encoding a myelin protein.
- 71. (New): The method of claim 70, wherein the myelin protein is myelin basic protein (MBP).
- 72. (New): The method of claim 69, further comprising a polynucleotide encoding an insulin protein.
- 73. (New): The method of claim 72, wherein the insulin protein is selected from the group consisting of insulin, proinsulin and preproinsulin.